

77. A dosage form as defined in claim 75, wherein said matrix comprises hydroxypropyl cellulose.
78. A dosage form as defined in claim 75, wherein said matrix comprises poly(ethylene oxide).
79. A dosage form as defined in claim 75, wherein said matrix comprises polyacrylic acid.
80. A dosage form as defined in claim 72, comprising a reservoir of azithromycin encased in a membrane which limits the release rate of azithromycin to said GI tract by diffusion.
81. A dosage form as defined in claim 80, in the form of a tablet coated with a membrane.
82. A dosage form as defined in claim 72, in the form of a multiparticulate comprising particles each of which is coated with a membrane which limits the release rate of said azithromycin by diffusion.
83. A dosage form as defined in claim 73, wherein a portion of the outside surface of said matrix is covered with an impermeable coating and the remainder of said outside surface is uncovered.
84. A dosage form as defined in claim 83, substantially in the shape of a cylinder wherein said impermeable coating covers one or both of the opposing flat surfaces thereof.
85. A dosage form as defined in claim 83, substantially in the shape of a cylinder wherein said impermeable coating covers only the radial surface thereof.
86. A dosage form as defined in claim 83, in the form of a tablet, wherein said uncovered area is in the form of an opening through said impermeable coating.
87. A dosage form as defined in claim 83, in the form of a tablet, wherein said uncovered area is in the form of a passageway which penetrates through the entire device.
88. A dosage form as defined in claim 83, in the form of a tablet, wherein said uncovered area is in the form of one or more slits through said impermeable coating or in the form of one or more strips removed therefrom.
89. A dosage form as defined in claim 83, substantially in the form of a cone, wherein the uncovered area is an opening for drug transport at or near the apex of the cone.
90. A dosage form as defined in claim 83, substantially in the shape of a hemisphere, wherein the uncovered area is in the form of an opening for drug transport at or near the center of the flat face of the hemisphere.
91. A dosage form as defined in claim 83, substantially in the shape of a half-cylinder, wherein the uncovered area is in the form of one or more slits at or near the centerline of the flat face of said half-cylinder.
92. A dosage form as defined in claim 72 in the form of a coated bi-layer tablet wherein one layer of said tablet comprises a water-swelling composition and the

002250-052200

second layer of said tablet comprises a dispensible azithromycin composition, said tablet being coated with a water-permeable membrane which is substantially impermeable to azithromycin, and which contains one or more perforations or passageways for exposing the azithromycin-containing composition to the use environment.

93. A dosage form as defined in claim 72 in the form of a coated tablet comprising a water-soluble salt of azithromycin, said tablet having a water-permeable coating which is substantially impermeable to azithromycin and substantially non-porous, said coating containing one or more perforations or passageways, for exposing the interior of the tablet to a use environment.

94. A dosage form as defined in claim 72 in the form of a coated tablet comprising azithromycin, said tablet having a porous coating which permits transport of both water and azithromycin through said porous coating.

95. A dosage form as defined in claim 72 in the form of a coated multiparticulate formulation wherein each particle comprises azithromycin and has a porous coating which permits transport of both water and azithromycin through said porous coating.

96. A delayed release dosage form of azithromycin which meets the following in vitro criteria:

in a first dissolution stage, $Q_{0.25} < 10\%$ when said dosage form is inserted in a USP rotating paddle apparatus, said apparatus being described in USP XXIII dissolution test chapter 711, and wherein said apparatus has paddles rotating at 50 rpm and contains 750 mL of 0.1 N HCl at 37°C;

in a second dissolution stage, $Q_{0.5} < Q_{0.25} + 10\%$ when 250 mL of 0.2 M tribasic sodium phosphate buffer is added to said acid immediately following said first stage to implement a buffer having a pH of about 6.8.

97. A dosage form as defined in claim 96, in the form of a tablet.

98. A dosage form as defined in claim 96, comprising a multiparticulate having a diameter between about 0.5 mm and about 3 mm.

99. A dosage form as defined in claim 96, comprising a multiparticulate having a diameter between about 0.1 and about 0.5 mm.

100. A dosage form as defined in claim 97, coated with a membrane comprising a polymer which is substantially insoluble and/or impermeable to azithromycin at the pH of the stomach, and is soluble and/or permeable to azithromycin at the pH of the small intestine and colon.

101. A dosage form as defined in claim 100, wherein said polymer is selected from cellulose acetate phthalate, polyvinylacetate phthalate, hydroxypropyl methylcellulose phthalate, and copolymers comprising acrylic acid and at least one acrylic acid ester.

102. A dosage form as defined in claim 98, wherein said multiparticulate is coated with a membrane comprising a polymer that is substantially insoluble and/or impermeable to azithromycin at the pH of the stomach, and is soluble and/or permeable to azithromycin at the pH of the small intestine and colon.

103. A dosage form as defined in claim 102, wherein said polymer is selected from cellulose acetate phthalate, polyvinylacetate phthalate, hydroxypropyl methylcellulose phthalate, and copolymers comprising acrylic acid and at least one acrylic acid ester.

Sub
C1
cont.

506
B2
002250-052200

Sub
C1
cont.

104. A dosage form as defined in claim 99, wherein said multiparticulate is coated with a membrane comprising a polymer that is substantially insoluble and/or impermeable to azithromycin at the pH of the stomach, and is soluble and/or permeable to azithromycin at the pH of the small intestine and colon.

105. A dosage form as defined in claim 104, wherein said polymer is selected from cellulose acetate phthalate, polyvinylacetate phthalate, hydroxypropyl methylcellulose phthalate, and copolymers comprising acrylic acid and at least one acrylic acid ester.

106. A dosage form as defined in claim 97, wherein said tablet optionally further comprises one or more osmagents, said tablet being surrounded by a semipermeable membrane that is permeable to water and substantially impermeable to said azithromycin and said osmagents.

107. A dosage form as defined in claim 98, said multiparticulate further comprising one or more osmagents, said multiparticulate being surrounded by a semipermeable membrane that is permeable to water and substantially impermeable to azithromycin and osmagents.

108. A dosage form as defined in claim 97, further comprising at least one swellable material, said tablet being surrounded by a semipermeable membrane that is permeable to water and substantially impermeable to azithromycin and said swellable material.

109. A dosage form as defined in claim 98, further comprising at least one swellable material, each multiparticulate particle being surrounded by a semipermeable membrane that is permeable to water and substantially impermeable to azithromycin and said swellable materials.

110. A dosage form as defined in claim 97, comprising:
a core comprising azithromycin and at least one osmagent;
a wall surrounding said tablet comprising a semipermeable membrane which is permeable to water and substantially impermeable to azithromycin and osmagent; and
a pH-sensitive trigger means attached to said semipermeable membrane for triggering the bursting of the tablet, said trigger means triggering at a pH between 3 and 9.

111. A dosage form as defined in claim 98, said multiparticulate further comprising one or more osmagents, each multiparticulate being surrounded by a wall comprising a semipermeable membrane which is permeable to water and substantially impermeable to azithromycin and osmagent; and
a pH-sensitive trigger means attached to said semipermeable membrane for triggering the bursting of the multiparticulate said trigger means triggering at pH between 3 and 9.

112. An azithromycin dosage form as defined in claim 110, wherein said core further comprises at least one swelling material.

113. An azithromycin dosage form as defined in claim 111, wherein said multiparticulate further each comprise at least one swelling material.

114. A dosage form as defined in claim 97, comprising:
a core comprising azithromycin and at least one osmagent;
a membrane surrounding said tablet core wherein said membrane is fabricated from a microporous hydrophobic support material;

Sub
C
Con

002250" 0504/5501

a hydrophobic liquid entrained within said membrane, said hydrophobic liquid being substantially impermeable to water and azithromycin, but being capable of changing to become substantially permeable to water and azithromycin.

115. A dosage form as defined in claim 98, comprising:
a core comprising azithromycin and at least one osmagent;
a membrane surrounding said multiparticulate core wherein said membrane is fabricated from a microporous hydrophobic support material;
a hydrophobic liquid entrained within said membrane, said hydrophobic liquid being substantially impermeable to water and azithromycin, but being capable of changing to become substantially permeable to water and azithromycin.

116. An azithromycin-containing dosage form as defined in claim 97, comprising:
a core comprising azithromycin and at least one swelling material;
a membrane surrounding said tablet core wherein said membrane is fabricated from a microporous hydrophobic support material;
a hydrophobic liquid entrained within said membrane, said hydrophobic liquid being substantially impermeable to water and azithromycin, but being capable of changing to become substantially permeable to water and azithromycin.

117. A dosage form as defined in claim 98, comprising:
a core comprising azithromycin and at least one swelling material;
a membrane surrounding said multiparticulate core wherein said membrane is fabricated from a microporous hydrophobic support material;
a hydrophobic liquid entrained within said membrane, said hydrophobic liquid being substantially impermeable to water and azithromycin, but being capable of changing to become substantially permeable to water and azithromycin.

118. A dosage form as defined in claim 97, comprising:
a core comprising azithromycin and at least one swelling material and/or at least one osmagent;
a membrane surrounding said tablet core wherein said membrane is substantially impermeable to azithromycin and labile to enzymes produced by bacteria which inhabit the colon.

119. A dosage form as defined in claim 98, comprising:
a core comprising azithromycin and at least one swelling material and/or at least one osmagent;
a membrane surrounding said multiparticulate core wherein said membrane is substantially impermeable to azithromycin and labile to enzymes produced by bacteria which inhabit the colon.

120. A dosage form as defined in claim 118, wherein said membrane comprises a polymer comprising at least one ethylenically unsaturated monomer crosslinked by a substituted or unsubstituted divinylazobenzene.

121. A dosage form as defined in claim 119, wherein said membrane comprises a polymer comprising at least one ethylenically unsaturated monomer crosslinked by a substituted or unsubstituted divinylazobenzene.

122. A dosage form as defined in claim 118, wherein said membrane comprises at least one polysaccharide.

123. A dosage form as defined in claim 119, wherein said membrane comprises at least one polysaccharide.

Sub
C
con

002250-052200

124. A dosage form as defined in claim 96, in the form of a capsule comprising two interpenetrating pieces, a first male piece comprising a water-swellaable material, which swells to effect disengagement of a second female piece upon administration to said mammal.

125. A sustained release dosage form of azithromycin for ingestion by a mammal which meets, based on the weight of said mammal, the following in vitro criteria:

- (1) $Q_{0.25} \leq 4$ mg/Kg of mammal weight,
- (2) $Q_1 \leq 10$ mg/Kg of mammal weight,
- (3) $Q_2 \leq 20$ mg/Kg of mammal weight,
- (4) $Q_4 \leq 30$ mg/Kg of mammal weight, and
- (5) $Q_6 \leq 40$ mg/Kg of mammal weight,

when said dosage form is tested in a USP rotating paddle apparatus, said apparatus being described in USP XXIII dissolution test chapter 711, and wherein the apparatus has paddles rotating at 50 rpm and contains 900 mL of pH 6.0 sodium dihydrogen phosphate buffer at 37°C;

and wherein, if said dosage form is a capsule, said buffer is implemented to contain 0.1 mg/mL of trypsin.

126. A dosage form as defined in claim 125, wherein said azithromycin is embedded in a matrix, which releases said azithromycin by diffusion.

127. A dosage form as defined in claim 126, wherein said matrix remains substantially intact during the period of drug release.

128. A dosage form as defined in claim 126, wherein said azithromycin is embedded in a matrix which releases said azithromycin by eroding.

129. A dosage form as defined in claim 128, wherein said matrix comprises hydroxypropyl methylcellulose.

130. A dosage form as defined in claim 128, wherein said matrix comprises hydroxypropyl cellulose.

131. A dosage form as defined in claim 128, wherein said matrix comprises poly(ethylene oxide).

132. A dosage form as defined in claim 128, wherein said matrix comprises polyacrylic acid.

133. A dosage form as defined in claim 125, comprising a reservoir of azithromycin encased in a membrane which limits the release rate of azithromycin to said GI tract by diffusion.

134. A dosage form as defined in claim 133, in the form of a tablet coated with a membrane.

135. A dosage form as defined in claim 135, in the form of a multiparticulate comprising particles each of which is coated with a membrane which limits the release rate of said azithromycin by diffusion.

136. A dosage form as defined in claim 126, wherein a portion of the outside surface of said matrix is covered with an impermeable coating and the remainder of said outside surface is uncovered.

137. A dosage form as defined in claim 136, substantially in the shape of a cylinder wherein said impermeable coating covers one or both of the opposing flat surfaces thereof.

138. A dosage form as defined in claim 136, substantially in the shape of a cylinder wherein said impermeable coating covers only the radial surface thereof.

139. A dosage form as defined in claim 136, in the form of a tablet, wherein said uncovered area is in the form of an opening through said impermeable coating.

140. A dosage form as defined in claim 136, in the form of a tablet, wherein said uncovered area is in the form of a passageway which penetrates through the entire device.

141. A dosage form as defined in claim 136, in the form of a tablet, wherein said uncovered area is in the form of one or more slits through said impermeable coating or in the form of one or more strips removed therefrom.

142. A dosage form as defined in claim 136, substantially in the form of a cone, wherein the uncovered area is an opening for drug transport at or near the apex of the cone.

143. A dosage form as defined in claim 136, substantially in the shape of a hemisphere, wherein the uncovered area is in the form of an opening for drug transport at or near the center of the flat face of the hemisphere.

144. A dosage form as defined in claim 136, substantially in the shape of a half-cylinder, wherein the uncovered area is in the form of one or more slits at or near the centerline of the flat face of said half-cylinder.

145. A dosage form as defined in claim 125 in the form of a coated bi-layer tablet, wherein one layer of said tablet comprises a water-swellable composition and the second layer of said tablet comprises a dispensible azithromycin composition, said tablet being coated with a water-permeable membrane which is substantially impermeable to azithromycin, and which contains one or more perforations or passageways for exposing the azithromycin-containing composition to the use environment.

146. A dosage form as defined in claim 125 in the form of a coated tablet comprising a water-soluble salt of azithromycin, said tablet having a water-permeable coating which is substantially impermeable to azithromycin and substantially non-porous, said coating containing one or more perforations or passageways, for exposing the interior of the tablet to a use environment.

147. A dosage form as defined in claim 125 in the form of a coated tablet comprising azithromycin, said tablet having a porous coating which permits transport of both water and azithromycin through said porous coating.

148. A dosage form as defined in claim 125 in the form of a coated multiparticulate formulation wherein each particle comprises azithromycin and has a porous coating which permits transport of both water and azithromycin through said porous coating.

Only after new claims 72-148 above have been entered, cancel original claim 1.

Sub
C
Cont.

with

with

with

with

with

with

002250-052200